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ORAL PRESENTATION

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Association of genomewide newborn DNA methylation patterns with maternal and newborn characteristics

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Background

Epigenetics, such as DNA methylation, plays a major role in development and disease risk. Work in model organisms clearly demonstrates that maternal, gestational, genetic and nutritional factors can greatly influence patterns of DNA methylation and increase the risk of disease. In collaboration with a large longitudinal cohort study of human development from gestation to age three, we examined how maternal age, maternal blood nutrient levels and genetic variation are related to neonatal genome-wide DNA methylation patterns.

Materials and methods

Genome-wide SNP variation was assayed in 96 mothers and their newborns and genome-wide patterns of DNA methylation in 120 newborns. In a subset of 30 mothers, maternal blood levels of several nutrients and metabolites in the one carbon pathway were directly measured. For DNA methylation, the unit of analysis was the average level of methylation across several CpGs within each of ~15,000 loci. Gene-by-gene, newborn DNA methylation was related to maternal age, maternal blood nutrient levels, and newborn SNP variation.

Results and conclusion

Genome-wide there is a generally negative correlation between maternal age and newborn methylation in CpG islands. At a subset of loci, particularly the X chromosome, there was a distinctly bimodal distribution of DNA methylation among newborns that was highly associated with SNP variants on other chromosomes.

Among maternal blood nutrients surveyed, phosphatidylcholine showed a significant correlation with newborn DNA methylation at a subset of loci.

Maternal age and blood nutrient levels are correlated with newborn DNA methylation patterns. Trans-acting SNPs are suggestively associated with bimodal patterns of DNA methylation at some loci.

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